

Sequence Listing

<110> Lee, Chichang; Ly, Celia; Moore, Gordon; Chi, Xiamei, Centocor, Inc.

<120> Methods and Compositions for Enhanced Protein Expression and/or Growth of Cultured Cells Using Co-Transcription of a Bcl2 Encoding Nucleic Acid

<130> CEN0269 PCT

<140> PCT/US01/45553

<141> 2001-11-02

<160> 14

<170> Patentin Ver 3.1

<210> 1

<211> 239

<212> PRT

<213> Homo sapiens

<400> 1

Met Ala His Ala Gly Arg Thr Gly Tyr Asp Asn Arg Glu lle Val Met

1

5

10

15

Lys Tyr Ile His Tyr Lys Leu Ser Gln Arg Gly Tyr Glu Trp Asp Ala

20

25

30

Gly Asp Val Gly Ala Ala Pro Pro Gly Ala Ala Pro Ala Pro Gly Ile

35

40

45

Phe Ser Ser Gln Pro Gly His Thr Pro His Pro Ala Ala Ser Arg Asp

50

55

60

Pro Val Ala Arg Thr Ser Pro Leu Gln Thr Pro Ala Ala Pro Gly Ala

65

70

75

Ala Ala Gly Pro Ala Leu Ser Pro Val Pro Pro Val Val His Leu Ala 85 90 95

Leu Arg Gln Ala Gly Asp Asp Phe Ser Arg Arg Tyr Arg Gly Asp Phe
100 105 110

Ala Glu Met Ser Ser Gln Leu His Leu Thr Pro Phe Thr Ala Arg Gly
115 120 125

Arg Phe Ala Thr Val Val Glu Glu Leu Phe Arg Asp Gly Val Asn Trp 130 135 140

Gly Arg Ile Val Ala Phe Phe Glu Phe Gly Gly Val Met Cys Val Glu 145 150 155 160

Ser Val Asn Arg Glu Met Ser Pro Leu Val Asp Asn Ile Ala Leu Trp 165 170 175

Met Thr Glu Tyr Leu Asn Arg His Leu His Thr Trp Ile Gln Asp Asn 180 185 190

Gly Gly Trp Asp Ala Phe Val Glu Leu Tyr Gly Pro Ser Met Arg Pro 195 200 205

Leu Phe Asp Phe Ser Trp Leu Ser Leu Lys Thr Leu Leu Ser Leu Ala 210 215 220

Leu Val Gly Ala Cys Ile Thr Leu Gly Ala Tyr Leu Ser His Lys 225 230 235

<210> 2

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<212> PRT

<213> Homo sapiens

<400> 2

Met Ala His Ala Gly Arg Thr Gly Tyr Asp Asn Arg Glu lle Val Met Lys Tyr lle His Tyr Lys Leu Ser Gln Arg Gly Tyr Glu Trp Asp Ala Gly Asp Val Gly Ala Ala Pro Pro Gly Ala Ala Pro Ala Pro Gly Ile Phe Ser Ser Gln Pro Gly His Thr Pro His Pro Ala Ala Ser Arg Asp Pro Val Ala Arg Thr Ser Pro Leu Gln Thr Pro Ala Ala Pro Gly Ala Ala Ala Gly Pro Ala Leu Ser Pro Val Pro Pro Val Val His Leu Ala Leu Arg Gln Ala Gly Asp Asp Phe Ser Arg Arg Tyr Arg Gly Asp Phe Ala Glu Met Ser Ser Gln Leu His Leu Thr Pro Phe Thr Ala Arg Gly Arg Phe Ala Thr Val Val Glu Glu Leu Phe Arg Asp Gly Val Asn Trp Gly Arg Ile Val Ala Phe Phe Glu Phe Gly Gly Val Met Cys Val Glu Ser Val Asn Arg Glu Met Ser Pro Leu Val Asp Asn Ile Ala Leu Trp Met Thr Glu Tyr Leu Asn Arg His Leu His Thr Trp lle Gln Asp Asn

Gly Gly Trp Val Gly Ala Ser Gly Asp Val Ser Leu Gly

200

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<210> 3

<211> 239

<212> PRT

<213> Homo sapiens

<400> 3

Met Ala His Ala Gly Arg Thr Gly Tyr Asp Asn Arg Glu lle Val Met

15

1 5 10

Lys Tyr Ile His Tyr Lys Leu Ser Gln Arg Gly Tyr Glu Trp Asp Ala

20 25 30

Gly Asp Val Gly Ala Ala Pro Pro Gly Ala Ala Pro Ala Pro Gly Ile

35 40 45

Phe Ser Ser Gln Pro Gly His Thr Pro His Pro Ala Ala Ser Arg Asp

50 55 60

Pro Val Ala Arg Thr Ser Pro Leu Gln Thr Pro Ala Ala Pro Gly Ala

65 70 75 80

Ala Ala Gly Pro Ala Leu Ser Pro Val Pro Pro Val Val His Leu Ala

85 90 95

Leu Arg Gln Ala Gly Asp Asp Phe Ser Arg Arg Tyr Arg Gly Asp Phe

100 105 110

Ala Glu Met Ser Ser Gln Leu His Leu Thr Pro Phe Thr Ala Arg Gly

115 120 125

Arg Phe Ala Thr Val Val Glu Glu Leu Phe Arg Asp Gly Val Asn Trp

130 135 140

Gly Arg Ile Val Ala Phe Phe Glu Phe Gly Val Met Cys Val Glu

145 150 155

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Ser Val Asn Arg Glu Met Ser Pro Leu Val Asp Asn Ile Ala Leu Trp
               170
                            175
                                         180
 Met Thr Glu Tyr Leu Asn Arg His Leu His Thr Trp Ile Gln Asp Asn
        180
                     185
                                   190
 Gly Gly Trp Asp Ala Phe Val Glu Leu Tyr Gly Pro Ser Met Arg Pro
      195
                   200
                                205
 Leu Phe Asp Phe Ser Trp Leu Ser Leu Lys Thr Leu Leu Ser Leu Ala
   210
                215
                              220
 Leu Val Gly Ala Cys Ile Thr Leu Gly Ala Tyr Leu Ser His Lys
 225
              230
                           235
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<213> Homo sapiens
<400> 4
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                      10
                                   15
Phe Tyr Phe Ala Ser
       20
<210> 5
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<213> Homo sapiens
<400> 5
Met Asp Glu Asp Val Leu Pro Gly Glu Val Leu Ala Ile Glu Gly Ile
         5
                     10
                                  15
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Phe Met Ala Cys Gly Leu Asn Glu Pro Glu Tyr Leu Tyr His Pro Leu 20 25 30

Leu Ser Pro Ile Lys Leu Tyr Ile Thr Gly Leu Met Arg Asp Lys Glu 35 40 45

Ser Leu Phe Glu Ala Met Leu Ala Asn Val Arg Phe His Ser Thr Thr 50 55 60

Gly Ile Asn Gln Leu Gly Leu Ser Met Leu Gln Val Ser Gly Asp Gly 65 70 75 180

Asn Met Asn Trp Gly Arg Ala Leu Ala IIe Leu Thr Phe Gly Ser Phe 85 90 95

Val Ala Gln Lys Leu Ser Asn Glu Pro His Leu Arg Asp Phe Ala Leu 100 105 110

Ala Val Leu Pro Val Tyr Ala Tyr Glu Ala Ile Gly Pro Gln Trp Phe 115 120 125

Arg Ala Arg Gly Gly Trp Arg Gly Leu Lys Ala Tyr Cys Thr Gln Val 130 135 140

Leu Thr Arg Arg Gly Arg Arg Met Thr Ala Leu Leu Gly Ser Ile 145 150 155 160

Ala Leu Leu Ala Thr lle Leu Ala Ala Val Ala Met Ser Arg Arg 165 170 175

<210> 6

<211> 211

<212> PRT

<213> Homo sapiens

<400> 6

Met Ala Ser Gly Gln Gly Pro Gly Pro Pro Arg Gln Ĝlu Cys Gly Glu Pro Ala Leu Pro Ser Ala Ser Glu Glu Gln Val Ala Gln Asp Thr Glu Glu Val Phe Arg Ser Tyr Val Phe Tyr Arg His Gln Gln Glu Gln Glu Ala Glu Gly Val Ala Ala Pro Ala Asp Pro Glu Met Val Thr Leu Pro Leu Gln Pro Ser Ser Thr Met Gly Gln Val Gly Arg Gln Leu Ala Ile Ile Gly Asp Asp Ile Asn Arg Arg Tyr Asp Ser Glu Phe Gln Thr Met Leu Gln His Leu Gln Pro Thr Ala Glu Asn Ala Tyr Glu Tyr Phe Thr Lys lle Ala Thr Ser Leu Phe Glu Ser Gly lle Asn Trp Gly Arg Val Val Ala Leu Leu Gly Phe Gly Tyr Arg Leu Ala Leu His Val Tyr Gln His Gly Leu Thr Gly Phe Leu Gly Gln Val Thr Arg Phe Val Val Asp Phe Met Leu His His Cys IIe Ala Arg Trp IIe Ala Gln Arg Gly Gly

Val Val Leu Gly Val Val Leu Leu Gly Gln Phe Val Val Arg Arg Phe

Trp Val Ala Ala Leu Asn Leu Gly Asn Gly Pro lie Leu Asn Val Leu

200

205

Phe Lys Ser

210

<210> 7

<211> 170

<212> PRT

<213> Homo sapiens

<400> 7

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1 5 10 15

Leu Ser Gln Lys Gly Tyr Ser Trp Ser Gln Phe Ser Asp Val Glu Glu

20 25 30

Asn Arg Thr Glu Ala Pro Glu Gly Thr Glu Ser Glu Met Glu Thr Pro

35 40 45

Ser Ala Ile Asn Gly Asn Pro Ser Trp His Leu Ala Asp Ser Pro Ala

50 55 60

Val Asn Gly Ala Thr Gly His Ser Ser Ser Leu Asp Ala Arg Glu Val

65 70 75 80

lle Pro Met Ala Ala Val Lys Gln Ala Leu Arg Glu Ala Gly Asp Glu

85 90 95

Phe Glu Leu Arg Tyr Arg Arg Ala Phe Ser Asp Leu Thr Ser Gln Leu

100 105 110

His Ile Thr Pro Gly Thr Ala Tyr Gln Ser Phe Glu Gln Asp Thr Phe

115 120 125

Val Glu Leu Tyr Gly Asn Asn Ala Ala Ala Glu Ser Arg Lys Gly Gln

Glu Arg Phe Asn Arg Trp Phe Leu Thr Gly Met Thr Val Ala Gly Val

Val Leu Leu Gly Ser Leu Phe Ser Arg Lys

<210> 8

<211> 160

<212> PRT

<213> Homo sapiens

<400> 8

Met Ser Glu Val Arg Pro Leu Ser Arg Asp Ile Leu Met Glu Thr Leu

Leu Tyr Glu Gln Leu Leu Glu Pro Pro Thr Met Glu Val Leu Gly Met

- 20

Thr Asp Ser Glu Glu Asp Leu Asp Pro Met Glu Asp Phe Asp Ser Leu

Glu Cys Met Glu Gly Ser Asp Ala Leu Ala Leu Arg Leu Ala Cys Ile

Gly Asp Glu Met Asp Val Ser Leu Arg Ala Pro Arg Leu Ala Gln Leu

Ser Glu Val Ala Met His Ser Leu Gly Leu Ala Phe Ile Tyr Asp Gln

Thr Glu Asp Ile Arg Asp Val Leu Arg Ser Phe Met Asp Gly Phe Thr

Thr Leu Lys Glu Asn lle Met Arg Phe Trp Arg Ser Pro Asn Pro Gly

Ser Trp Val Ser Cys Glu Gln Val Leu Leu Ala Leu Leu Leu Leu 130 135 140

Ala Leu Leu Leu Pro Leu Leu Ser Gly Gly Leu His Leu Leu Leu Lys 145 150 155 160

<210> 9

<211> 218

<212> PRT

<213> Homo sapiens

<400> 9

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1 5 10 15

Glu Gin Ile Met Lys Thr Gly Ala Leu Leu Leu Gin Gly Phe Ile Gin 20 25 30

Asp Arg Ala Gly Arg Met Gly Gly Glu Ala Pro Glu Leu Ala Leu Asp 35 40 45

Pro Val Pro Gln Asp Ala Ser Thr Lys Lys Leu Ser Glu Cys Leu Lys 50 55 60

Arg Ile Gly Asp Glu Leu Asp Ser Asn Met Glu Leu Gln Arg Met Ile 65 70 75 80

Ala Ala Val Asp Thr Asp Ser Pro Arg Glu Val Phe Phe Arg Val Ala 85 90 95

Ala Asp Met Phe Ser Asp Gly Asn Phe Asn Trp Gly Arg Val Val Ala 100 105 110

Leu Phe Tyr Phe Ala Ser Lys Leu Val Leu Lys Ala Leu Cys Thr Lys 115 120 125

Val Pro Glu Leu IIe Arg Thr IIe Met Gly Trp Thr Leu Asp Phe Leu Arg Glu Arg Leu Leu Gly Trp Ile Gln Asp Gln Gly Gly Trp Val Arg Leu Leu Lys Pro Pro His Pro His His Arg Ala Leu Thr Thr Ala Pro Ala Pro Pro Ser Leu Pro Pro Ala Thr Pro Leu Gly Pro Trp Ala Phe Trp Ser Arg Ser Gln Trp Cys Pro Leu Pro lle Phe Arg Ser Ser Asp Val Val Tyr Asn Ala Phe Ser Leu Arg Val <210> 10 <211> 239 <212> PRT <213> Homo sapiens <400> 10 Met Ala His Ala Gly Arg Thr Gly Tyr Asp Asn Arg Glu Ile Val Met Lys Tyr Ile His Tyr Lys Leu Ser Gln Arg Gly Tyr Glu Trp Asp Ala Gly Asp Val Gly Ala Ala Pro Pro Gly Ala Ala Pro Ala Pro Gly lle Phe Ser Ser Gln Pro Gly His Thr Pro His Pro Ala Ala Ser Arg Asp

Pro Val Ala Arg Thr Ser Pro Leu Gin Thr Pro Ala Ala Pro Gly Ala

Ala Ala Gly Pro Ala Leu Ser Pro Val Pro Pro Val Val His Leu Thr

Leu Arg Gin Ala Gly Asp Asp Phe Ser Arg Arg Tyr Arg Arg Asp Phe

Ala Glu Met Ser Ser Gln Leu His Leu Thr Pro Phe Thr Ala Arg Gly

Arg Phe Ala Thr Val Val Glu Glu Leu Phe Arg Asp Gly Val Asn Trp

Gly Arg Ile Val Ala Phe Phe Glu Phe Gly Gly Val Met Cys Val Glu

Ser Val Asn Arg Glu Met Ser Pro Leu Val Asp Asn Ile Ala Leu Trp

Met Thr Glu Tyr Leu Asn Arg His Leu His Thr Trp Ile Gln Asp Asn

Gly Gly Trp Asp Ala Phe Val Glu Leu Tyr Gly Pro Ser Met Arg Pro

Leu Phe Asp Phe Ser Trp Leu Ser Leu Lys Thr Leu Leu Ser Leu Ala

Leu Val Gly Ala Cys lle Thr Leu Gly Ala Tyr Leu Gly His Lys

<210> 11

<211> 205

<212> PRT

<213> Homo sapiens

<400> 11
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1 5 10 15

Lys Tyr lle His Tyr Lys Leu Ser Gln Arg Gly Tyr Glu Trp Asp Ala 20 25 30

Gly Asp Val Gly Ala Ala Pro Pro Gly Ala Ala Pro Ala Pro Gly Ile 35 40 45

Phe Ser Ser Gln Pro Gly His Thr Pro His Pro Ala Ala Ser Arg Asp 50 55 60

Pro Val Ala Arg Thr Ser Pro Leu Gln Thr Pro Ala Ala Pro Gly Ala 65 70 75 80

Ala Ala Gly Pro Ala Leu Ser Pro Val Pro Pro Val Val His Leu Ala 85 90 95

Leu Arg Gln Ala Gly Asp Asp Phe Ser Arg Arg Tyr Arg Gly Asp Phe 100 105 110

Ala Glu Met Ser Ser Gln Leu His Leu Thr Pro Phe Thr Ala Arg Gly
115 120 125

Arg Phe Ala Thr Val Val Glu Glu Leu Phe Arg Asp Gly Val Asn Trp 130 135 140

Gly Arg Ile Val Ala Phe Phe Glu Phe Gly Gly Val Met Cys Val Glu 145 150 155 160

Ser Val Asn Arg Glu Met Ser Pro Leu Val Asp Asn Ile Ala Leu Trp 165 170 175

Met Thr Glu Tyr Leu Asn Arg His Leu His Thr Trp Ile Gln Asp Asn 180 185 190

Gly Gly Trp Val Gly Ala Ser Gly Asp Val Ser Leu Gly <210> 12 <211> 245 <212> PRT <213> Homo sapiens <400> 12 Leu Ala Gin Arg Giy Giy Ala Arg Arg Pro Arg Giy Asp Arg Giu Arg Leu Gly Ser Arg Leu Arg Ala Leu Arg Pro Gly Arg Glu Pro Arg Gln Ser Glu Pro Pro Ala Gln Arg Gly Pro Pro Pro Ser Arg Arg Pro Pro Ala Arg Ser Thr Ala Ser Gly His Asp Arg Pro Thr Arg Gly Ala Ala Ala Gly Ala Arg Arg Pro Arg Met Lys Lys Lys Thr Arg Arg Arg Ser Thr Arg Ser Glu Glu Leu Thr Arg Ser Glu Glu Leu Thr Leu Ser Glu Glu Ala Thr Trp Ser Glu Glu Ala Thr Gln Ser Glu Glu Ala Thr Gln Gly Glu Glu Met Asn Arg Ser Gln Glu Val Thr Arg Asp Glu Glu Ser Thr Arg Ser Glu Glu Val Thr Arg Glu Glu Met Ala Ala Ala Gly Leu

Thr Val Thr Val Thr His Ser Asn Glu Lys His Asp Leu His Val Thr

145 150 155 160

Ser Gln Gln Gly Ser Ser Glu Pro Val Val Gln Asp Leu Ala Gln Val 165 170 175

Val Glu Val Ile Gly Val Pro Gln Ser Phe Gln Lys Leu Ile Phe 180 185 190

Lys Gly Lys Ser Leu Lys Glu Met Glu Thr Pro Leu Ser Ala Leu Gly 195 200 205

Ile Gin Asp Gly Cys Arg Val Met Leu Ile Gly Lys Lys Asn Ser Pro 210 215 220

Gln Glu Glu Val Glu Leu Lys Lys Leu Lys His Leu Glu Lys Ser Val 225 230 235 240

Glu Lys lle Ala Asp Gln Leu Glu Glu Leu Asn Lys Glu Leu Thr Gly 245 250 255

Ile Gin Gin Giy Phe Leu Pro Lys Asp Leu Gin Ala Giu Ala Leu Cys 260 265 270

Lys Leu Asp Arg Arg Val Lys Ala Thr Ile Glu Gln Phe Met Lys Ile 275 280 285

Leu Glu Glu Ile Asp Thr Leu Ile Leu Pro Glu Asn Phe Lys Asp Ser 290 295 300

Arg Leu Lys Arg Lys Gly Leu Val Lys Lys Val Gln Ala Phe Leu Ala 305 310 315 320

Glu Cys Asp Thr Val Glu Gln Asn lle Cys Gln Glu Thr Glu Arg Leu 325 330 335

Gln Ser Thr Asn Phe Ala Leu Ala Glu 340 345

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<400> 13
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Lys Tyr lle His Tyr Lys Leu Ser Gln Arg Gly Tyr Glu Trp Asp Ala
       20
                    25
                                  30
Gly Asp Val Gly Ala Ala Pro Pro Gly Ala Ala Pro Ala Pro Gly Ile
     35
                  40
Phe Ser Ser Gln Pro Gly His Thr Pro His Pro Ala Ala Ser Arg Asp
   50
               55
                             60
Pro Val Ala Arg Thr Ser Pro Leu Gln Thr Pro Ala Ala Pro Gly Ala
65
             70
                          75
                                       80
Ala Ala Gly Pro Ala Leu Ser Pro Val Pro Pro Val Val His Leu Thr
         85
                      90
                                    95
Leu Arg Gln Ala Gly Asp Asp Phe Ser Arg Arg Tyr Arg Arg Asp Phe
       100
                    105
                                  110
Ala Glu Met Ser Ser Gln Leu His Leu Thr Pro Phe Thr Ala Arg Gly
     115
                  120
                                125
Arg Phe Ala Thr Val Val Glu Glu Leu Phe Arg Asp Gly Val Asn Trp
  130
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Gly Arg lie Val Ala Phe Phe Glu Phe Gly Val Met Cys Val Glu

Ser Val Asn Arg Glu Met Ser Pro Leu Val Asp Asn Ile Ala Leu Trp 170 175 Met Thr Glu Tyr Leu Asn Arg His Leu His Thr Trp Ile Gln Asp Asn 180 185 190 Gly Gly Trp Asp Ala Phe Val Glu Leu Tyr Gly Pro Ser Met Arg Pro 195 200 205 Leu Phe Asp Phe Ser Trp Leu Ser Leu Lys Thr Leu Leu Ser Leu Ala 210 215 220 Leu Val Gly Ala Cys lle Thr Leu Gly Ala Tyr Leu Gly His Lys 225 230 235 <210> 14 <211> 195 <212> PRT <213> Homo sapiens <400> 14 Met Asp Cys Glu Val Asn Asn Gly Ser Ser Leu Arg Asp Glu Cys Ile 1 5 10 15

Phe Arg Arg Glu Leu Asp Ala Leu Gly His Glu Leu Pro Val Leu Ala 35 40 45

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Pro Gln Trp Glu Gly Tyr Asp Glu Leu Gln Thr Asp Gly Asn Arg Ser 50 55 60

Ser His Ser Arg Leu Gly Arg Ile Glu Ala Asp Ser Glu Ser Gln Glu 65 70 75 80 Asp lle lle Arg Asn lle Ala Arg His Leu Ala Gln Val Gly Asp Ser 85 90 95

Met Asp Arg Ser IIe Pro Pro Gly Leu Val Asn Gly Leu Ala Leu Gln 100 105 110

Leu Arg Asn Thr Ser Arg Ser Glu Glu Asp Arg Asn Arg Asp Leu Ala 115 120 125

Thr Ala Leu Glu Gln Leu Leu Gln Ala Tyr Pro Arg Asp Met Glu Lys 130 135 140

Glu Lys Thr Met Leu Val Leu Ala Leu Leu Leu Ala Lys Lys Val Ala 145 150 155 160

Ser His Thr Pro Ser Leu Leu Arg Asp Val Phe His Thr Thr Val Asn 165 170 175

Phe Ile Asn Gln Asn Leu Arg Thr Tyr Val Arg Ser Leu Ala Arg Asn 180 185 190

Gly Met Asp 195

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